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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/815,242	03/21/2001	Robert Haselbeck	ELITRA.011A	7191
210 MERCK P O BOX 2000 RAHWAY, NJ 07065-0907	7590 11/04/2010		EXAMINER SCHNIZER, RICHARD A	
			ART UNIT 1635	PAPER NUMBER
			MAIL DATE 11/04/2010	DELIVERY MODE PAPER

Please find below and/or attached an Office communication concerning this application or proceeding.

The time period for reply, if any, is set in the attached communication.

Office Action Summary	Application No. 09/815,242	Applicant(s) HASELBECK ET AL.	
	Examiner Richard Schnizer	Art Unit 1635	

-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --

Period for Reply

A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) OR THIRTY (30) DAYS, WHICHEVER IS LONGER, FROM THE MAILING DATE OF THIS COMMUNICATION.

- Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication.
- If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication.
- Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).

Status

- 1) ☒ Responsive to communication(s) filed on 28 September 2010.
- 2a) ☒ This action is **FINAL**. 2b) ☐ This action is non-final.
- 3) ☐ Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under *Ex parte Quayle*, 1935 C.D. 11, 453 O.G. 213.

Disposition of Claims

- 4) ☒ Claim(s) 12,31,45-69,77-87,89-96,100,101,103 and 104 is/are pending in the application.
- 4a) Of the above claim(s) _____ is/are withdrawn from consideration.
- 5) ☐ Claim(s) _____ is/are allowed.
- 6) ☒ Claim(s) 12,31,45-69,77-86,89-96,100,101 and 103 is/are rejected.
- 7) ☐ Claim(s) 87 and 104 is/are objected to.
- 8) ☐ Claim(s) _____ are subject to restriction and/or election requirement.

Application Papers

- 9) ☒ The specification is objected to by the Examiner.
- 10) ☒ The drawing(s) filed on 21 March 2001 is/are: a) ☒ accepted or b) ☐ objected to by the Examiner.
Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).
Replacement drawing sheet(s) including the correction is required if the drawing(s) is objected to. See 37 CFR 1.121(d).
- 11) ☐ The oath or declaration is objected to by the Examiner. Note the attached Office Action or form PTO-152.

Priority under 35 U.S.C. § 119

- 12) ☐ Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).
- a) ☐ All b) ☐ Some * c) ☐ None of:
1. ☐ Certified copies of the priority documents have been received.
2. ☐ Certified copies of the priority documents have been received in Application No. _____.
3. ☐ Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).

* See the attached detailed Office action for a list of the certified copies not received.

Attachment(s)

- | | |
|---|---|
| 1) <input type="checkbox"/> Notice of References Cited (PTO-892) | 4) <input type="checkbox"/> Interview Summary (PTO-413) |
| 2) <input type="checkbox"/> Notice of Draftperson's Patent Drawing Review (PTO-948) | Paper No(s)/Mail Date. _____ |
| 3) <input type="checkbox"/> Information Disclosure Statement(s) (PTO/SB/08) | 5) <input type="checkbox"/> Notice of Informal Patent Application |
| Paper No(s)/Mail Date _____ | 6) <input type="checkbox"/> Other: _____ |

DETAILED ACTION

An amendment was received and entered on 9/28/10.

Claims 12, 31, 45-69, 77-87, 89-96, 100, 101, 103, and 104 remain pending and under consideration. The elected species of invention under consideration is a method as set forth in independent claims 12, 31, and 100, wherein the species of prokaryotic organism is *S. aureus* (Applicant's response of 6/30/03).

Election/Restrictions

Applicant's arguments filed 9/22/09 are persuasive with regard to SEQ ID NOS: 8502 and 5283, and these sequences are rejoined. Applicant also argues that SEQ ID NOS: 1390, 1845, 2782, and 3283 should also be examined because they are part of the larger molecule represented by SEQ ID NO: 4228 (elected) and so should not provide too great a burden. This is unpersuasive. Due to the extremely large number of hits obtained when searching SEQ ID NO: 4228, an effective search of these specific sequences requires a separate search query of multiple databases for each SEQ ID. As explained previously in the Actions of 8/22/02, 6/3/03, and 9/23/03, these searches present an undue burden on the Office due to the complex nature of the searches and the amount of processor time taken by each search.

Specification

The disclosure stands objected to because it contains an embedded hyperlink and/or other form of browser-executable code, e.g. at page 5, lines 18 and 19; page 76,

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line 11; page 90, lines 1, 4, 12, 14, 15, 17, and 18; and page 185, line 1. Applicant is required to delete all embedded hyperlinks and/or other forms of browser-executable code. See MPEP § 608.01.

In the response filed 9/28/10, Applicant directed amendments to specific paragraphs of US Publication 2002/0061569 that correspond to the paragraphs objected to above. The Examiner is unable to enter such amendments. Amendments to specifications, subject to entry by the Examiner, are made under 37 CFR 1.121. This regulation does not address amendments to US Patent Application Publications. It is suggested that Applicant direct amendments to the specification of the instant application under 37 CFR 1.121.

Claim Rejections - 35 USC § 102

The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless –

(b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.

Claims 12, 31, 45-52, 57, 85, 86, 89-92, and 101 stand rejected under 35 U.S.C. 102(b) as being anticipated by Vermuelen et al (US 5872104) as evidenced by Schaefer et al (J. Bact. 188(23): 8252-8258, 2006).

Vermuelen taught a method of sensitizing bacteria to the effects of an antibiotic by inhibiting the expression of bacterial methylases through the use of antisense, and then contacting the sensitized bacteria with an antibiotic. See abstract; Example II at

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columns 32 and 33; and Example XII at columns 67-70. In particular, Example II describes how different combinations of sensitizing agents and antibiotics are screened. In one test, no sensitizing compound is added (sentence bridging columns 32 and 33. Organisms in which this method can be performed include *S. aureus* and coagulase negative staphylococci (see entire document , e.g. Table 6 column 29; and Table 7 at column 30).

Note that the instant claims require a sublethal level of an antisense nucleic acid comprising "a nucleotide sequence of SEQ ID NO: 1463. Thus any sublethal level of any antisense nucleic acid comprising any sequence of SEQ ID NO: 1463, as few as two nucleotides, can be used in the methods as claimed.

SEQ ID NO: 1463 inhibits the expression of *S. aureus* yphC (Applicant's response of 9/22/09 at page 15) which encodes a GTPase required for the assembly of the large ribosomal subunit (see Schaefer (2006)). Thus the expected effect of treating a prokaryotic cell with SEQ ID NO: 1463 would be to decrease the expression of yphC protein, resulting in inhibition of ribosome synthesis, and a decrease in expression and activity of all protein gene products. In other words, the expression, and subsequently the activity, of all *S. aureus* protein gene products, including the methylases of Vermuelen, is inhibited by SEQ ID 1463. So, Vermuelen taught a method of screening for antibiotics by providing to an *S. aureus*, or coagulase negative staphylococcus, varying non-lethal amounts of an antisense that inhibits the expression or activity of a methylase enzyme. This produces sensitized bacteria. The sensitized bacteria are contacted with a variety of antibiotics to determine which combination of antisense and

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antibiotic kills bacteria most efficiently. Because the methylase enzyme must be produced by ribosome-driven translation, it is a gene product whose expression and activity are inhibited by SEQ ID NO: 1463 because the gene product inhibited by SEQ ID NO: 1463 is required for translation of any and all *S. aureus* proteins.

Thus Vermuelen teaches each and every limitation of the invention and anticipates the claims.

Response to Arguments

Applicant's arguments filed 9/28/10 have been fully considered but they are not persuasive.

At page 24 of the response, Applicant argues that claim 12 requires an antisense nucleic acid comprising SEQ ID NO: 1463. This is incorrect. Claim 12 requires only that the antisense nucleic acid comprise a sequence of SEQ ID NO: 1463. Thus any antisense comprising any two nucleotide sequence present in SEQ ID NO: 1463 (in other words, any antisense nucleic acid at all) meets the claim requirements.

Applicant also argues that claim 31 requires that the claimed method decrease the activity or amount of a gene product encoded by a nucleic acid that hybridizes to SEQ ID NO: 1463. This is incorrect. Claim 31 allows claimed method to decrease the activity or amount of a gene product encoded by a nucleic acid comprising a nucleotide sequence which hybridizes to a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 1463. See item "ii)". Thus the activity or amount of any gene product can be decreased. This is because "a nucleotide sequence of SEQ ID NO: 1463" means any

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dinucleotide present in SEQ ID NO: 1463. Since SEQ ID NO: 1463 contains all possible dinucleotides there is essentially no limitation on the gene product whose activity or amount is decreased.

Applicant's arguments related to claim 100 are irrelevant because claim 100 was not rejected over Vermuelen. Note that claim 101, which was rejected, depends from claims 48 and 31.

Claim Rejections - 35 USC § 103

The following is a quotation of 35 U.S.C. 103(a) which forms the basis for all obviousness rejections set forth in this Office action:

(a) A patent may not be obtained though the invention is not identically disclosed or described as set forth in section 102 of this title, if the differences between the subject matter sought to be patented and the prior art are such that the subject matter as a whole would have been obvious at the time the invention was made to a person having ordinary skill in the art to which said subject matter pertains. Patentability shall not be negated by the manner in which the invention was made.

Claim 12, 31, 45-69, 77-86, 89-96, 100, and 101 stand rejected under 35 U.S.C. 103(a) as being unpatentable over Zalacain et al (WO 01/23418), Fritz et al (US 6627747), Zhang et al (Gene 255: 297-305, 2000), and Ji et al (J. Bact. 181(21): 6585-6590, 1999).

Zalacain disclosed yphC gene and polypeptide from *S. aureus*, and suggested that yphC could be used as a target for the development of antibiotics (page 2, lines 5-12; and page 16, line 10 to page 22, line 9). Zalacain also showed that yphC was essential for growth in *S. aureus* by allelic replacement in strain RN4220 (page 32, lines 15-30). The Zalacain publication designates the United States and claims priority to Non-Provisional Application 09/406,968, filed 9/28/1999. The information relating to

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Zalacain that is used in this rejection is completely supported by the '968 application and is entitled to a filing date of 9/28/1999.

Fritz taught that yphC was an essential GTPase in *S. pneumoniae*, that the gene was conserved and essential in other bacteria, such as *B. subtilis* and *E. coli*, and suggested screening for inhibitors of yphC because such inhibitors would have a broad spectrum of antibiotic activity. See abstract; paragraph bridging columns 13 and 14; column 16, lines 34-50; and column 19, lines 49-60. Fritz suggested testing candidate YphC inhibitors to determine if they can inhibit bacterial growth (see column 26, lines 33 to 47 and column 28, lines 41-57).

These references did not teach a method of screening for antibiotics in which bacterial cells were sensitized to the effects of a potential antibiotic by decreasing the amount or activity of yphC.

Zhang taught methods and systems for determining gene essentiality in bacteria, and for determining if the mode of action of an antibiotic involves a given essential gene. The methods rely on systems that allow one to modulate the level of expression of a known or suspected essential gene. Regulatable expression systems were developed that exhibit titratable induction of expression. By using these systems to vary the amount of expression of an essential gene in a bacterium such as *S. aureus* or *Bacillus subtilis*, and treating the bacterium with a candidate antibiotic, one can determine if any growth inhibition caused by the antibiotic is due to a mode of action involving the essential gene. This can be done by varying the amount of expression of a particular essential gene in separate cultures, and then determining the minimum

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inhibitory concentration of a candidate antibiotic in each culture. If the function of an antibiotic involves the targeted essential gene, then the sensitivity of the cells to the antibiotic will vary inversely with the amount of the gene that is expressed. See abstract and section 3.3 on pages 302-304. Cell growth rates were measured by monitoring optical density at 600 nm (section 2.7 on page 300). Strains used included *S. aureus* RN4220 (section 2.1 on page 298).

It would have been obvious to one of ordinary skill in the art at the time of the invention to screen for yphC inhibitors because both Zalacain and Fritz taught that yphC was a target for antibiotic development. One would have been motivated to use the method of Zhang because it allows one to determine if the cellular activity of an antibiotic is occurring through its proposed molecular target as opposed to some other mechanism. Moreover, yphC was known to be essential in *S. aureus*, *S. pneumoniae*, *B. subtilis*, and *E. coli* (see Zalacain at page 32, lines 15-30; and Fritz at columns 13 and 14; column 16, lines 34-50; and column 19, lines 49-60) such that inhibitors that act through yphC are expected to be broad spectrum antibiotics.

This method does not involve the use of antisense, as does the instantly claimed method.

Ji taught a method and system for regulated expression of antisense in *S. aureus*, and showed that gene expression could be efficiently inhibited by expressing antisense in situ in *S. aureus* RN4220. See abstract.

In view of the teachings of Zhang it was clear that one could determine if a candidate antibiotic was acting through its intended target by varying the amount of

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expression of that target in the presence of the antibiotic. It was also clear at the time of the invention that there was more than one way to modulate the expression of an essential gene. Zhang taught a method in which gene expression was controlled by replacing the chromosomal copy of the essential gene with another copy under the control of an inducible promoter. On the other hand Ji taught a method of regulating expression of essential genes in *S. aureus* by expressing antisense RNA directed against the transcript of the essential gene. Thus it would have been clear to one of ordinary skill in the art at the time of the invention that one could sensitize cells for screening antibiotics by regulating the expression of *yphC* negatively, i.e. by using antisense to control the expression of the essential gene, as taught by Ji, instead of using positive control of gene expression, as taught by Zhang. This is simply a matter of design choice. All of the technology required to control gene expression, either positively or negatively, was known in the art, and one of ordinary skill could have chosen either approach and achieved the same end, i.e. control over the amount of *yphC* expressed in the subject *S. aureus* culture. In using the antisense approach, it would have been obvious to refer to the prior art sequence of Zalacain to obtain an antisense sequence. Instant SEQ ID NO: 1463 is 100% complementary to nucleotides 122-508 of the *yphC* sequence of Zalacain, see alignment below. Accordingly, the antisense obtained would necessarily downregulate an RNA that can be reduced by antisense comprising a sequence of SEQ ID NO: 1463, as required by claim 12 and dependents.

Score = 715 bits (387), Expect = 0.0
Identities = 387/387 (100%), Gaps = 0/387 (0%)
Strand=Plus/Minus

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"Query" = yphC DNA sequence from Zalacain

"Subject" = complement of instant SEQ IDNO: 1463

Query	122	GTATTTATTCTTCAGGTGAATGGTTAACACATGATTTCAATATTATTGATACAGGTGGTA	181
Sbjct	387	GTATTTATTCTTCAGGTGAATGGTTAACACATGATTTCAATATTATTGATACAGGTGGTA	328
Query	182	TTGAAATTGGTGATGCACCATTCCAAACACAAATTAGAGCGCAGGCAGAAATCGCCATAG	241
Sbjct	327	TTGAAATTGGTGATGCACCATTCCAAACACAAATTAGAGCGCAGGCAGAAATCGCCATAG	268
Query	242	ATGAAGCGGATGTTATTATTTTTATGGTTAACGTGCGTGAAGGATTGACACAAAGCGATG	301
Sbjct	267	ATGAAGCGGATGTTATTATTTTTATGGTTAACGTGCGTGAAGGATTGACACAAAGCGATG	208
Query	302	AAATGGTCGCTCAAATTTTATACAAATCTAAAAAACCGGTCGTATTAGCGGTTAACAAAG	361
Sbjct	207	AAATGGTCGCTCAAATTTTATACAAATCTAAAAAACCGGTCGTATTAGCGGTTAACAAAG	148
Query	362	TAGATAATATGGAAATGCGTACAGACGTGTATGATTTCTATTATTAGGATTTGGTGAAC	421
Sbjct	147	TAGATAATATGGAAATGCGTACAGACGTGTATGATTTCTATTATTAGGATTTGGTGAAC	88
Query	422	CGTATCCGATATCAGGGTCACATGGTTTAGGTCTTGGTGACTTGTTAGATGCAGTTGTTT	481
Sbjct	87	CGTATCCGATATCAGGGTCACATGGTTTAGGTCTTGGTGACTTGTTAGATGCAGTTGTTT	28
Query	482	CTCATTTTGGTGAAGAGGAAGAAGATC	508
Sbjct	27	CTCATTTTGGTGAAGAGGAAGAAGATC	1

Moreover, the polypeptide sequence of Zalacain is 100% identical to instant SEQ ID

NO: 12600 (see alignment below), which is encoded by instant SEQ ID NO: 8502. Thus

one would have inhibited the expression of instant SEQ IDNO: 12600 as intended in

instant claims 31, 100, and dependents.

Score = 887 bits (2293), Expect = 0.0, Method: Compositional matrix adjust.

Identities = 436/436 (100%), Positives = 436/436 (100%), Gaps = 0/436 (0%)

"Query" = yphC amino acid sequence from Zalacain

"Subject" = instant SEQ ID NO: 12600

Query	1	MTKPIVAIVGRPNVGKSTIFNRIVGERVSIVEDTPGVTRDRIYSSGEWLTHDFNIIDTGG	60
		MTKPIVAIVGRPNVGKSTIFNRIVGERVSIVEDTPGVTRDRIYSSGEWLTHDFNIIDTGG	
Sbjct	1	MTKPIVAIVGRPNVGKSTIFNRIVGERVSIVEDTPGVTRDRIYSSGEWLTHDFNIIDTGG	60
Query	61	IEIGDAPFQTQIRAQAEIAIDEADVIIIFMVNVREGLTQSDMVAQILYKSKKPVVLAVNK	120
		IEIGDAPFQTQIRAQAEIAIDEADVIIIFMVNVREGLTQSDMVAQILYKSKKPVVLAVNK	
Sbjct	61	IEIGDAPFQTQIRAQAEIAIDEADVIIIFMVNVREGLTQSDMVAQILYKSKKPVVLAVNK	120

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Query	121	VDNMEMRTDVYDFYSLGFGEYPYISGSHGLGLGDLDDAVVSHFGEEEDPYDEDTIRLSI	180
		VDNMEMRTDVYDFYSLGFGEYPYISGSHGLGLGDLDDAVVSHFGEEEDPYDEDTIRLSI	
Sbjct	121	VDNMEMRTDVYDFYSLGFGEYPYISGSHGLGLGDLDDAVVSHFGEEEDPYDEDTIRLSI	180
Query	181	IGRPNVGKSSLVNAILGEDRVIVSNVAGTTRDAIDTEYSYDGQDYVLIDTAGMRKKGKVY	240
		IGRPNVGKSSLVNAILGEDRVIVSNVAGTTRDAIDTEYSYDGQDYVLIDTAGMRKKGKVY	
Sbjct	181	IGRPNVGKSSLVNAILGEDRVIVSNVAGTTRDAIDTEYSYDGQDYVLIDTAGMRKKGKVY	240
Query	241	ESTEKYSVLRALKAIERSNVVLVVIDAEQGIIEQDKRVAGYAHEQGKAVVIVVNKWDTVE	300
		ESTEKYSVLRALKAIERSNVVLVVIDAEQGIIEQDKRVAGYAHEQGKAVVIVVNKWDTVE	
Sbjct	241	ESTEKYSVLRALKAIERSNVVLVVIDAEQGIIEQDKRVAGYAHEQGKAVVIVVNKWDTVE	300
Query	301	KDSKTMKKFEDEVKRFQFLDYAQIAFVSAKERTRLRTLFPYINEASENHKKRVQSSTLN	360
		KDSKTMKKFEDEVKRFQFLDYAQIAFVSAKERTRLRTLFPYINEASENHKKRVQSSTLN	
Sbjct	301	KDSKTMKKFEDEVKRFQFLDYAQIAFVSAKERTRLRTLFPYINEASENHKKRVQSSTLN	360
Query	361	EVVTDAISMNPTPTDKGRRLNVFYATQVAIEPPTFVVVFVNDVELMHFSYKRYLENQIRAA	420
		EVVTDAISMNPTPTDKGRRLNVFYATQVAIEPPTFVVVFVNDVELMHFSYKRYLENQIRAA	
Sbjct	361	EVVTDAISMNPTPTDKGRRLNVFYATQVAIEPPTFVVVFVNDVELMHFSYKRYLENQIRAA	420
Query	421	FGFEGTPIHIIARKRN	436
		FGFEGTPIHIIARKRN	
Sbjct	421	FGFEGTPIHIIARKRN	436

Claims 78-84 require that the antisense nucleic acid used to reduce the amount or activity of the recited gene product must comprise a sequence with at least 70-97% nucleotide sequence identity to SEQ ID NO: 1463. These claims are included in the rejection because it would have been obvious to one of ordinary skill in the art at the time of the invention to use the entire coding sequence of yphC to generate an antisense transcript, i.e. to use an antisense complementary to the entire length of the yphC transcript. In so doing one would obtain an antisense transcript comprising SEQ ID NO: 1463. Moreover, the claims as written do not specify the length of the sequence that must share identity with SEQ ID NO: 1463, such that an antisense molecule with as few as two nucleotides of SEQ ID NO: 1463 "comprises a sequence having at least 97% nucleotide sequence identity to SEQ ID NO: 1463." SEQ ID NO: 1463 contains all 16 possible dinucleotide sequences, so any antisense directed to yphC will contain at

least one dinucleotide that is 100% identical to SEQ ID NO: 1463. Because the cited references render obvious antisense directed to the yphC sequence of Zalacain, they render obvious an antisense nucleic acid as required by instant claims 78-84.

Thus the invention as a whole was prima facie obvious.

Response to Arguments

Applicant's arguments filed 9/28/10 have been fully considered but they are not persuasive.

Applicant argues that the cited references do not teach the claimed invention. Applicant engages in a discussion of supposed disadvantages of traditional methods of drug discovery, and advantages of the use of sensitized cells, however Applicant fails to identify any specific claim limitation that is not taught by the cited references as combined. It is noted that, contrary to Applicant's assertions regarding sensitized cells and screening in vivo, at pages 25 and 26, Zhang taught the use of sensitized cells to screen in vivo. More particularly, Zhang showed that by varying the amount of expression of an essential gene in a bacterium such as *S. aureus* or *Bacillus subtilis*, and treating the bacterium with a candidate antibiotic, one can determine if any growth inhibition caused by the antibiotic is due to a mode of action involving the essential gene, thereby addressing Applicant's arguments regarding the "noise" of compounds acting at targets that are not of interest". See section .33 bridging pages 302-304, wherein Zhang discusses the use of the system to screen one of several candidate

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antibiotics to determine if it acts through the essential gene encoding polypeptide deformylase.

Applicant argues that neither Zalacain nor Fritz shows any way of finding compounds targeted to yphC that have an effect in vivo. This is unpersuasive because Zalacain and Fritz were not relied on to teach in vivo screening. Zalacain and Fritz disclosed yphC as an essential gene, and suggested screening for inhibitors of its activity. Zhang provided a means for screening for inhibitors that act through a mechanism involving an essential gene of interest. One of ordinary skill aware of these disclosures would clearly take advantage of the method of Zhang to screen candidate inhibitors, as suggested by Zalacain and Fritz, because the method allows one to detect inhibitors that act through a mechanism involving the essential gene of interest, i.e. it is a means of identifying relevant inhibitors. Moreover, it would have been obvious to modify the method of Zhang to manipulate the amount of expression of an essential gene of interest using antisense for the reasons set forth in the rejection.

Conclusion

No claim is allowed. Claims 87 and 104 stand objected to because they recite non-elected subject matter and depend from rejected claims, but would be allowable if rewritten in independent form excluding the nonelected subject matter and incorporating all of the limitations of the claims from which they depend.

THIS ACTION IS MADE FINAL. Applicant is reminded of the extension of time policy as set forth in 37 CFR 1.136(a).

A shortened statutory period for reply to this final action is set to expire THREE MONTHS from the mailing date of this action. In the event a first reply is filed within TWO MONTHS of the mailing date of this final action and the advisory action is not mailed until after the end of the THREE-MONTH shortened statutory period, then the shortened statutory period will expire on the date the advisory action is mailed, and any extension fee pursuant to 37 CFR 1.136(a) will be calculated from the mailing date of the advisory action. In no event, however, will the statutory period for reply expire later than SIX MONTHS from the date of this final action.

Any inquiry concerning this communication or earlier communications from the examiner(s) should be directed to Richard Schnizer, whose telephone number is 571-272-0762. The examiner can normally be reached Monday through Friday between the hours of 6:00 AM and 3:30 PM. The examiner is off on alternate Fridays, but is sometimes in the office anyway.

If attempts to reach the examiner by telephone are unsuccessful, the Examiner's acting supervisor, Heather Calamita, can be reached at (571) 272-2876. The official central fax number is 571-273-8300. Any inquiry of a general nature or relating to the status of this application or proceeding should be directed to (571) 272-0547.

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Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see <http://pair-direct.uspto.gov>. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

/Richard Schnizer/
Primary Examiner, Art Unit 1635